

## ENTERED

PCT10

RAW SEQUENCE LISTING

PATENT APPLICATION: US/10/049,182

DATE: 02/25/2002

TIME: 10:48:27

Input Set : A:\36667.txt

```
4 <110> APPLICANT: Banks, William A.
      6 <120> TITLE OF INVENTION: MODULATION OF THE BLOOD-BRAIN BARRIER TRANSPORTER FOR
             LEPTIN
     7
     9 <130> FILE REFERENCE: 01017/35040
C--> 11 <140> CURRENT APPLICATION NUMBER: US/10/049,182
C--> 12 <141> CURRENT FILING DATE: 2002-02-08
     14 <160> NUMBER OF SEQ ID NOS: 6
     16 <170> SOFTWARE: PatentIn Ver. 2.0
     18 <210> SEQ ID NO: 1
     19 <211> LENGTH: 2793
     20 <212> TYPE: DNA
     21 <213> ORGANISM: Murine
     23 <220> FEATURE:
     24 <223> OTHER INFORMATION: Murine ob (leptin) cDNA
     26 <220> FEATURE:
     27 <221> NAME/KEY: CDS
     28 <222> LOCATION: (57)..(557)
     30 <220> FEATURE:
     31 <221> NAME/KEY: sig_peptide
     32 <222> LOCATION: (57)..(59)
     34 <220> FEATURE:
     35 <221> NAME/KEY: mat_peptide
     36 <222> LOCATION: (60)..(557)
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     40
     41
     43 tgc tgg aga ccc ctg tgt cgg ttc ctg tgg ctt tgg tcc tat ctg tct
                                                                           107
     44 Cys Trp Arg Pro Leu Cys Arg Phe Leu Trp Leu Trp Ser Tyr Leu Ser
                                              10
     47 tat gtt caa gca gtg cct atc cag aaa gtc cag gat gac acc aaa acc
     48 Tyr Val Gln Ala Val Pro Ile Gln Lys Val Gln Asp Asp Thr Lys Thr
                                                              30
                     20
                                          25
     51 ctc atc aag acc att gtc acc agg atc aat gac att tca cac acg cag
                                                                            203
     52 Leu Ile Lys Thr Ile Val Thr Arg Ile Asn Asp Ile Ser His Thr Gln
                                      40
                                                          45
     53
                                                                            251
     56 tcg gta tcc gcc aag cag agg gtc act ggc ttg gac ttc att cct ggg
     57 Ser Val Ser Ala Lys Gln Arg Val Thr Gly Leu Asp Phe Ile Pro Gly
                                  55
                                                                            299
     60 ctt cac ccc att ctg agt ttg tcc aag atg gac cag act ctg gca gtc
     61 Leu His Pro Ile Leu Ser Leu Ser Lys Met Asp Gln Thr Leu Ala Val
                                                                       80
                             70
                                                  75
     62 65
     64 tat caa cag gtc ctc acc agc ctg cct tcc caa aat gtg ctg cag ata
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| 65 Tyr Gln Gln Val Leu Thr Ser Leu Pro Ser Gln Asn Val Leu Gln Ile<br>66 85 90 95   |  |  |  |  |  |  |  |  |  |  |  |
|---|--|--|--|--|--|--|--|--|--|--|--|
| 68 gcc aat gac ctg gag aat ctc cga gac ctc ctc cat ctg ctg gcc ttc 395  |  |  |  |  |  |  |  |  |  |  |  |
| 69 Ala Asn Asp Leu Glu Asn Leu Arg Asp Leu Leu His Leu Leu Ala Phe  |  |  |  |  |  |  |  |  |  |  |  |
| 70 100 105 110  |  |  |  |  |  |  |  |  |  |  |  |
| 72 tcc aag agc tgc tcc ctg cct cag acc agt ggc ctg cag aag cca gag 443  |  |  |  |  |  |  |  |  |  |  |  |
| 73 Ser Lys Ser Cys Ser Leu Pro Gln Thr Ser Gly Leu Gln Lys Pro Glu  |  |  |  |  |  |  |  |  |  |  |  |
| 74 115 120 125  |  |  |  |  |  |  |  |  |  |  |  |
| 76 ago ctg gat ggo gto ctg gaa goo toa cto tac too aca gag gtg gtg 491  |  |  |  |  |  |  |  |  |  |  |  |
| 77 Ser Leu Asp Gly Val Leu Glu Ala Ser Leu Tyr Ser Thr Glu Val Val 78 130 135 140   |  |  |  |  |  |  |  |  |  |  |  |
| 78 130 135 140<br>80 gct ttg agc agg ctg cag ggc tct ctg cag gac att ctt caa cag ttg 539  |  |  |  |  |  |  |  |  |  |  |  |
| 81 Ala Leu Ser Arg Leu Gln Gly Ser Leu Gln Asp Ile Leu Gln Gln Leu  |  |  |  |  |  |  |  |  |  |  |  |
| 82 145 150 155 160  |  |  |  |  |  |  |  |  |  |  |  |
| 84 gat gtt agc cct gaa tgc tgaagtttca aaggccacca ggctcccaag 587   |  |  |  |  |  |  |  |  |  |  |  |
| 85 Asp Val Ser Pro Glu Cys  |  |  |  |  |  |  |  |  |  |  |  |
| 86 165  |  |  |  |  |  |  |  |  |  |  |  |
| 88 aatcatgtag agggaagaaa ccttggcttc caggggtctt caggagaaga gagccatgtg 647  |  |  |  |  |  |  |  |  |  |  |  |
| 90 cacacateca teatteattt eteteeetee tgtagaceae ceatecaaag geatgactee 707  |  |  |  |  |  |  |  |  |  |  |  |
| 92 acaatgettg acteaagtta tecacacaac tteatgagea caaggagggg ceageetgea 767  |  |  |  |  |  |  |  |  |  |  |  |
| 94 gaggggactc tcacctagtt cttcagcaag tagagataag agccatccca tcccctccat 827  |  |  |  |  |  |  |  |  |  |  |  |
| 96 gtcccacctg ctccgggtac atgttcctcc gtgggtacac gcttcgctgc ggcccaggag 887  |  |  |  |  |  |  |  |  |  |  |  |
| 98 aggtgaggta gggatgggta gagcctttgg gctgtctcag agtctttggg agcaccgtga 947  |  |  |  |  |  |  |  |  |  |  |  |
| 100 aggetgeate cacacacage tggaaactee caageageae acgatggaag cacttattta 1007  |  |  |  |  |  |  |  |  |  |  |  |
| 102 tttattctgc attctatttt ggatggatct gaagcaaggc atcagctttt tcaggctttg 1067  |  |  |  |  |  |  |  |  |  |  |  |
| 104 ggggtcagcc aggatgagga aggctcctgg ggtgctgctt tcaatcctat tgatgggtct 1127 106 gcccgaggca aacctaattt ttgagtgact ggaaggaagg ttgggatctt ccaaacaaga 1187     |  |  |  |  |  |  |  |  |  |  |  |
| 108 gtctatgcag gtagcgctca agattgacct ctggtgactg gttttgtttc tattgtgact 1247  |  |  |  |  |  |  |  |  |  |  |  |
| 110 gactetatee aaacaegttt geageggeat tgeegggage ataggetagg ttattateaa 1307  |  |  |  |  |  |  |  |  |  |  |  |
| 112 aagcagatga attttgtcaa gtgtaatatg tatctatgtg cacctgaggg tagaggatgt 1367  |  |  |  |  |  |  |  |  |  |  |  |
| 114 gttagaggga gggtgaagga tccggaagtg ttctctgaat tacatatgtg tggtaggctt 1427  |  |  |  |  |  |  |  |  |  |  |  |
| 116 ttctgaaagg gtgaggcatt ttcttacctc tgtggccaca tagtgtggct ttgtgaaaag 1487  |  |  |  |  |  |  |  |  |  |  |  |
| 118 gacaaaggag ttgactcttt ccggaacatt tggagtgtac caggcaccct tggaggggct 1547  |  |  |  |  |  |  |  |  |  |  |  |
| 120 aaagctacag gccttttgtt ggcatattgc tgagctcagg gagtgagggc cccacatttg 1607  |  |  |  |  |  |  |  |  |  |  |  |
| 122 agacagtgag ccccaagaaa agggtccctg gtgtagatct ccaaggttgt ccagggttga 1667  |  |  |  |  |  |  |  |  |  |  |  |
| 124 teteacaatg egittettaa geaggiagae gittgeatge eaatatgigg tieteatetg 1727  |  |  |  |  |  |  |  |  |  |  |  |
| 126 attggttcat ccaaagtaga accetgtcte ccaeccatte tgtggggagt tttgttccag 1787  |  |  |  |  |  |  |  |  |  |  |  |
| 128 tgggaatgag aaatcactta gcagatggte etgagecetg ggceageact getgaggaag 1847  |  |  |  |  |  |  |  |  |  |  |  |
| 130 tgccagggcc ccaggccagg ctgccagaat tgcccttcgg gctggaggat gaacaaaggg 1907 132 gcttgggttt ttccatcacc cctgcaccct atgtcaccat caaactgggg ggcagatcag 1967     |  |  |  |  |  |  |  |  |  |  |  |
| 132 gettigggttt tideateade detgeadeet atgicadeat caaactyggg gycagateag 1507<br>134 tgagaggaca cttgatggaa agcaatacac tttaagactg agcacagttt cgtgctcagc 2027 |  |  |  |  |  |  |  |  |  |  |  |
| 136 tetgtetggt getgtgaget agagaagete accaeataea tataaaaate agaggeteat 2087  |  |  |  |  |  |  |  |  |  |  |  |
| 138 gtccctgtgg ttagacccta ctcgcggcgg tgtactccac cacagcagca ccgcaccgct 2147  |  |  |  |  |  |  |  |  |  |  |  |
| 140 ggaagtacag tgctgtcttc aacaggtgtg aaagaacctg agctgagggt gacagtgccc 2207  |  |  |  |  |  |  |  |  |  |  |  |
| 142 aggggaaccc tgcttgcagt ctattgcatt tacataccgc atttcagggc acattagcat 2267  |  |  |  |  |  |  |  |  |  |  |  |
| 144 ccactcctat ggtagcacac tgttgacaat aggacaaggg ataggggttg actatcctt 2327   |  |  |  |  |  |  |  |  |  |  |  |
| 146 atccaaaatg cttgggacta gaagagtttt ggattttaga gtcttttcag gcataggtat 2387  |  |  |  |  |  |  |  |  |  |  |  |
| 148 atttgagtat atataaaatg agatatcttg gggatggggc ccaagtataa acatgaagtt 2447  |  |  |  |  |  |  |  |  |  |  |  |
| 150 catttatatt tcataatacc gtatagacac tgcttgaagt gtagttttat acagtgtttt 2507  |  |  |  |  |  |  |  |  |  |  |  |
|   |  |  |  |  |  |  |  |  |  |  |  |

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152 aaataacgtt gtatgcatga aagacgtttt tacagcatga acctgtctac tcatgccagc 2567
154 actcaaaaac cttggggttt tggagcagtt tggatcttgg gttttctgtt aagagatggt 2627
156 tagettatae etaaaaceat aatggeaaae aggetgeagg accagaetgg atceteagee 2687
158 ctgaagtgtg cccttccagc caggtcatac cctgtggagg tgagcgggat caggttttgt 2747
160 ggtgctaaga gaggagttgg aggtagattt tggaggatct gagggc
163 <210> SEQ ID NO: 2
164 <211> LENGTH: 167
165 <212> TYPE: PRT
166 <213> ORGANISM: Murine
168 <400> SEQUENCE: 2
169 Met Cys Trp Arg Pro Leu Cys Arg Phe Leu Trp Leu Trp Ser Tyr Leu
                          5
                                             10
170 -1 1
172 Ser Tyr Val Gln Ala Val Pro Ile Gln Lys Val Gln Asp Asp Thr Lys
                                         25
175 Thr Leu Ile Lys Thr Ile Val Thr Arg Ile Asn Asp Ile Ser His Thr
176
                 35
178 Gln Ser Val Ser Ala Lys Gln Arg Val Thr Gly Leu Asp Phe Ile Pro
181 Gly Leu His Pro Ile Leu Ser Leu Ser Lys Met Asp Gln Thr Leu Ala
                             70
         65
184 Val Tyr Gln Gln Val Leu Thr Ser Leu Pro Ser Gln Asn Val Leu Gln
                         85
                                              90
187 Ile Ala Asn Asp Leu Glu Asn Leu Arg Asp Leu Leu His Leu Leu Ala
                                         105
                    100
190 Phe Ser Lys Ser Cys Ser Leu Pro Gln Thr Ser Gly Leu Gln Lys Pro
                                    120
                115
191
193 Glu Ser Leu Asp Gly Val Leu Glu Ala Ser Leu Tyr Ser Thr Glu Val
                                135
            130
196 Val Ala Leu Ser Arg Leu Gln Gly Ser Leu Gln Asp Ile Leu Gln Gln
                            150
                                                 155
199 Leu Asp Val Ser Pro Glu Cys
200 160
                        165
203 <210> SEQ ID NO: 3
204 <211> LENGTH: 700
205 <212> TYPE: DNA
206 <213> ORGANISM: Homo sapiens
208 <220> FEATURE:
209 <221> NAME/KEY: CDS
210 <222> LOCATION: (46)..(546)
212 <220> FEATURE:
213 <221> NAME/KEY: sig_peptide
214 <222> LOCATION: (46)..(48)
216 <220> FEATURE:
217 <221> NAME/KEY: mat_peptide
218 <222> LOCATION: (49)..(546)
220 <220> FEATURE:
221 <223> OTHER INFORMATION: Human ob (leptin) where N represents adenine or
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224 <400> SEQUENCE: 3
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RAW SEQUENCE LISTING DATE: 02/25/2002 PATENT APPLICATION: US/10/049,182 TIME: 10:48:27

Input Set : A:\36667.txt

| W> | 225<br>226<br>227               | nnng  | gnngt    | tg ( | caagg     | jecea      | ia ga | n gaageeeann nteetgggaa g |     |           |           |     |     | ggaaa atg cat tgg gga 57<br>Met His Trp Gly<br>-1 1 |           |                   |            |     |  |
|----|---------------------------------|---|----------|------|-----------|------------|-------|---------------------------|-----|-----------|-----------|-----|-----|---|-----------|-------------------|------------|-----|--|
|    | 229                             |   |          |      |           |            |       |                           |     |           |           |     |     | ttc   | tat       | gtc<br>Val        |            | 105 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | atc<br>Ile        |            | 153 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | gtc<br>Val<br>50  |            | 201 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | cac<br>His        |            | 249 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | caa<br>Gln        |            | 297 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | aac<br>Asn        |            | 345 |  |
|    |                                 | Leu   |          |      |           |            |       |                           |     |           |           |     |     |   |           | aag<br>Lys        |            | 393 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | ctg<br>Leu<br>130 |            | 441 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | ctg<br>Leu        |            | 489 |  |
|    |                                 |   |          |      |           |            |       |                           |     |           |           |     |     |   |           | ctc<br>Leu        |            | 537 |  |
| W> |                                 | 69 cct ggg tgc tgaggccttg aaggtcactc ttcctgcaag gactnacgtt 70 Pro Gly Cys 71 165  |          |      |           |            |       |                           |     |           |           |     |     |   |           |                   | 586        |     |  |
|    | 272<br>274<br>277<br>278<br>279 | 2 aagggaagga actetggttt ceaggtatet eeaggattga agageattge atggacaeee<br>24 ettateeagg actetgteaa ttteeetgae teetetaage eactetteea aagg<br>27 <210> SEQ ID NO: 4<br>28 <211> LENGTH: 167<br>29 <212> TYPE: PRT<br>30 <213> ORGANISM: Homo sapiens |          |      |           |            |       |                           |     |           |           |     |     |   |           | acaccc            | 646<br>700 |     |  |
|    | 283<br>284<br>286               | Met<br>-1   | His<br>1 | Trp  | Gly       | Thr<br>Ala | 5     |                           |     |           | Lys       | 10  |     |   |           | Tyr               | 15         |     |  |
|    | 287<br>289<br>290               | Thr   | Leu      | Ile  | Lys<br>35 | 20<br>Thr  | Ile   | Val                       | Thr | Arg<br>40 | 25<br>Ile | Asn | Asp | Ile   | Ser<br>45 | 30<br>His         | Thr        |     |  |

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```
292 Gln Ser Val Ser Ser Lys Gln Lys Val Thr Gly Leu Asp Phe Ile Pro
           50
                              55
295 Gly Leu His Pro Ile Leu Thr Leu Ser Lys Met Asp Gln Thr Leu Ala
                          70
298 Val Tyr Gln Gln Ile Leu Thr Ser Met Pro Ser Arg Asn Val Ile Gln
                                         90
                      8.5
301 Ile Ser Asn Asp Leu Glu Asn Leu Arg Asp Leu Leu His Val Leu Ala
                                    105
                  100
304 Phe Ser Lys Ser Cys His Leu Pro Trp Ala Ser Gly Leu Glu Thr Leu
305 115
                                 120
307 Asp Ser Leu Gly Gly Val Leu Glu Ala Ser Gly Tyr Ser Thr Glu Val
308 130
                             135
310 Val Ala Leu Ser Arg Leu Gln Gly Ser Leu Gln Asp Met Leu Trp Gln
                         150
311 145
313 Leu Asp Leu Ser Pro Gly Cys
314 160
317 <210> SEQ ID NO: 5
318 <211> LENGTH: 146
319 <212> TYPE: PRT
320 <213> ORGANISM: Mus musculus
322 <220> FEATURE:
323 <223> OTHER INFORMATION: Mature mouse ob (leptin)
325 <400> SEQUENCE: 5
326 Val Pro Ile Gln Lys Val Gln Asp Asp Thr Lys Thr Leu Ile Lys Thr
327 1 5
                                      10
329 Ile Val Thr Arq Ile Asn Asp Ile Ser His Thr Gln Ser Val Ser Ala
               20
332 Lys Gln Arg Val Thr Gly Leu Asp Phe Ile Pro Gly Leu His Pro Ile
                              40
335 Leu Ser Leu Ser Lys Met Asp Gln Thr Leu Ala Val Tyr Gln Gln Val
336 50
                          55
338 Leu Thr Ser Leu Pro Ser Gln Asn Val Leu Gln Ile Ala Asn Asp Leu
                      70
                                          75
341 Glu Asn Leu Arg Asp Leu Leu His Leu Leu Ala Phe Ser Lys Ser Cys
                                      90
                   85
344 Ser Leu Pro Gln Thr Ser Gly Leu Gln Lys Pro Glu Ser Leu Asp Gly
345
              100
                                 105
347 Val Leu Glu Ala Ser Leu Tyr Ser Thr Glu Val Val Ala Leu Ser Arg
                             120
350 Leu Gln Gly Ser Leu Gln Asp Ile Leu Gln Gln Leu Asp Val Ser Pro
351 130
                         135
353 Glu Cys
354 145
357 <210> SEQ ID NO: 6
358 <211> LENGTH: 146
359 <212> TYPE: PRT
360 <213> ORGANISM: Homo sapiens
362 <220> FEATURE:
363 <223> OTHER INFORMATION: Mature human ob (leptin)
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VERIFICATION SUMMARY

PATENT APPLICATION: US/10/049,182

DATE: 02/25/2002

TIME: 10:48:28

Input Set : A:\36667.txt

Output Set: N:\CRF3\02252002\J049182.raw

L:11 M:270 C: Current Application Number differs, Replaced Application Number

L:12 M:271 C: Current Filing Date differs, Replaced Current Filing Date

L:225 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3 L:269 M:341 W: (46) "n" or "Xaa" used, for SEQ ID#:3